IB2004/003352

A. CLASSIFICATION OF SUBJECT MATTER IPC 7 A61K38/17

According to International Patent Classification (IPC) or to both national classification and IPC

#### **B. FIELDS SEARCHED**

Minimum documentation searched (classification system followed by classification symbols)

IPC 7 A61K C12N

Documentation searched other than minimum documentation to the extent that such documents are included in the fields searched

Electronic data base consulted during the international search (name of data base and, where practical, search terms used)

EPO-Internal, WPI Data, PAJ, BIOSIS, EMBASE, Sequence Search

Category °	Citation of document, with indication, where appropriate, of the relevant passages	Relevant to daim No.
X	TEGETHOFF S ET AL: "Tetrameric oligomerization of IkappaB kinase gamma (IKKgamma) is obligatory for IKK complex activity and NF-kappaB activation."  MOLECULAR AND CELLULAR BIOLOGY, vol. 23, no. 6, March 2003 (2003-03), pages 2029-2041, XP002314348 ISSN: 0270-7306 abstract, results; discussion;  -/	1-8

Y Further documents are listed in the continuation of box C.	Patent family members are listed in annex.			
<ul> <li>Special categories of cited documents:</li> <li>"A" document defining the general state of the art which is not considered to be of particular relevance</li> <li>"E" earlier document but published on or after the International filing date</li> <li>"L" document which may throw doubts on priority claim(s) or which is cited to establish the publication date of another citation or other special reason (as specified)</li> <li>"O" document referring to an oral disclosure, use, exhibition or other means</li> <li>"P" document published prior to the international filing date but later than the priority date claimed</li> </ul>	<ul> <li>"T" later document published after the international filing date or priority date and not in conflict with the application but cited to understand the principle or theory underlying the invention</li> <li>"X" document of particular relevance; the claimed invention cannot be considered novel or cannot be considered to involve an inventive step when the document is taken alone</li> <li>"Y" document of particular relevance; the claimed invention cannot be considered to involve an inventive step when the document is combined with one or more other such documents, such combination being obvious to a person skilled in the art.</li> <li>"&amp;" document member of the same patent family</li> </ul>			
Date of the actual completion of the international search	Date of mailing of the international search report			
20 January 2005	14/02/2005			
Name and mailing address of the ISA  European Palent Office, P.B. 5818 Patentlaan 2  NL – 2280 HV Rijswijk  Tel. (+31–70) 340–2040, Tx. 31 651 epo ni,	Authorized officer			
Fax: (+31-70) 340-3016	Sommer, B			

IB2004/003352

	ation) DOCUMENTS CONSIDERED TO BE RELEVANT	Rejevent to claim No.
Category °	Citation of document, with indication, where appropriate, of the relevant passages	Relevant to claim No.
X	LE PAGE C ET AL: "Disruption of NF-kappaB signaling and chemokine gene activation by retroviral mediated expression of IKKgamma/NEMO mutants" VIROLOGY, vol. 286, no. 2, 1 August 2001 (2001-08-01), pages 422-433, XP002314349 ISSN: 0042-6822 abstract; results; discussion;	1-8
X	WO 99/47672 A (WALLACH DAVID ; YEDA RES & DEV (IL); KOVALENKO ANDREI (IL); EINSTEIN C) 23 September 1999 (1999-09-23)	1-8
Y	page 10, line 3 - page 23, line 5; page 49, line 22 - page 50, line 6; claims; figures 3, 11, 12; example 10;	9–32
X	WO 02/092761 A (FANSLOW WILLIAM C III; IMMUNEX CORP (US); DERRY JONATHAN M J (US); DO) 21 November 2002 (2002-11-21) claims	1-8
X	WO 01/83554 A (PRAECIS PHARM INC; MAY MICHAEL J (US); UNIV YALE (US); FINDEIS MARK A) 8 November 2001 (2001-11-08)	1-8, 33-37
Υ	page 4, paragraph 3; page 15, paragraph 2 - page 16, paragraph 4; page 23, paragraph 3; claims; figures 2, 5; example 1;	9-32, 38-45
<b>X</b>	MAY M J ET AL: "Selective inhibition of NF-kB activation by a peptide that blocks the interaction of NEMO with the IkB kinase complex" SCIENCE, vol. 289, 1 September 2000 (2000-09-01), pages 1550-1554, XP002189523	1-8, 33-37
Y	ISSN: 0036-8075	9-32, 38-45
P,X .	AGOU F ET AL: "The trimerization domain of nemo is composed of the interacting C-terminal CC2 and LZ coiled-coil subdomains" JOURNAL OF BIOLOGICAL CHEMISTRY, vol. 279, no. 27, 2 July 2004 (2004-07-02), pages 27861-27869, XP002314350 ISSN: 0021-9258 the whole document	1-45
	-/	

[B2004/003352

Category °	Citation of document, with indication, where appropriate, of the relevant passages	Relevant to claim No.
Ρ,Χ	IHA H ET AL: "Segregation of NF-kappaB activation through NEMO/IKKgamma by Tax and TNFalpha: Implications for stimulus-specific interruption of oncogenic signaling."  ONCOGENE, vol. 22, no. 55, 4 December 2003 (2003-12-04), pages 8912-8923, XP002314351 ISSN: 0950-9232 abstract; page 8916, left-hand column, paragraph 3 - page 8919, right-hand column, paragraph 1; discussion, in particular page 8921, left-hand column, last paragraph; figures 4-7;	1-45
Т	AGOU F ET AL: "Inhibition of NF-kB activation by peptides targeting NF-kB essential modulator (NEMO) oligomerization" JOURNAL OF BIOLOGICAL CHEMISTRY, vol. 279, no. 52, 24 December 2004 (2004-12-24), pages 54248-54257, XP002314352 ISSN: 0021-9258	1-45

PCT/IB2004/003352

Box II Observations where certain claims were found unsearchable (Continuation of item 2 of first sheet)
This International Search Report has not been established in respect of certain claims under Article 17(2)(a) for the following reasons:
1. Claims Nos.: because they relate to subject matter not required to be searched by this Authority, namely:
Claims Nos.:     because they relate to parts of the International Application that do not comply with the prescribed requirements to such an extent that no meaningful International Search can be carried out, specifically:
3. Claims Nos.: because they are dependent claims and are not drafted in accordance with the second and third sentences of Rule 6.4(a).
Box III Observations where unity of invention is lacking (Continuation of item 3 of first sheet)
This International Searching Authority found multiple inventions in this international application, as follows:
see additional sheet
·
As all required additional search fees were timely paid by the applicant, this International Search Report covers all searchable claims.
2. X As all searchable claims could be searched without effort justifying an additional fee, this Authority did not invite payment of any additional fee.
3. As only some of the required additional search fees were timely paid by the applicant, this International Search Report covers only those claims for which fees were paid, specifically claims Nos.:
4. No required additional search fees were timely paid by the applicant. Consequently, this International Search Report is restricted to the Invention first mentioned in the claims; it is covered by claims Nos.:
Remark on Protest  The additional search fees were accompanied by the applicant's protest.  No protest accompanied the payment of additional search fees.

## FURTHER INFORMATION CONTINUED FROM PCT/ISA/ 210

This International Searching Authority found multiple (groups of) inventions in this international application, as follows:

1. claims: 1-32 (all partially)

a purified polynucleotide which encodes a polypeptide that inhibits the NFkappaB signaling pathway, said polypeptide comprising an amino acid sequence selected from SEQ ID Nos. 2, 3, 13 or 14, i.e. comprising the CC2 domain of NEMO, as well as subject-matter related thereto;

2. claims: 1-32 (all partially)

a purified polynucleotide which encodes a polypeptide that inhibits the NFkappaB signaling pathway, said polypeptide comprising an amino acid sequence selected from SEQ ID Nos. 6, 7, 15 or 16, i.e. comprising the LZ domain of NEMO, as well as subject-matter related thereto;

3. claims: 1-32 (all partially)

a purified polynucleotide which encodes a polypeptide that inhibits the NFkappaB signaling pathway, said polypeptide comprising an amino acid sequence selected from SEQ ID Nos. 30-39, i.e. comprising the mutated NEMO-like motif (NLM) domain of NEMO, as well as subject-matter related thereto;

4. claims: 33-45

a method of identifying polypeptides that modulate oligomerization of NEMO comprising a) identifying a candidate polypeptide sequence, b) linking said candidate polypeptide sequence to a polypeptide having a high transduction potential via a spacer sequence, c) contacting a cell culture with the polypeptide fusion construct, d) monitoring the activity of the NFkappaB signaling pathway, e) determining the relative inhibition by said polypeptide fusion construct and f) correlating relative inhibition by said polypeptide fusion construct to NEMO oligomerization as well as subject-matter related thereto;

IB2004/003352

	atent document d in search report		Publication date		Patent family member(s)		Publication date
WO	9947672		23~09-1999	AU	760900	B2	22-05-2003
				AU	2954599	A	11-10-1999
				BG	104769	Α	30-04-2001
				BR	9909659	Α	21-11-2000
				CA	2323637	A1	23-09-1999
				CN	1295614		16-05-2001
				EΑ	4062		25-12-2003
	•			EE	200000538		15-04-2002
				ΕP	1454985		08-09-2004
				EP	1062336		27-12-2000
				HU	0101612		28-01-2004
				MO	9947672		23-09-1999
				JP	2002506644		05-03-2002
			•	NO	20004649		30-10-2000
				NZ	506776		29-08-2003
				NZ	525566		24-12-2004
				PL	343262		30-07-2001
				SK	13762000		12-03-2001
	•			US	6734174		11-05-2004
				US ZA	2004219615 200004756		04-11-2004 11-01-2001
WO	02092761	Α	21-11-2002	US	2003165985		04-09-2003
				WO	02092761	A2	21-11-2002
WO	0183554	Α	08-11-2001	ΑT	279438		15-10-2004
				AU	5763101		12-11-2001
				Αυ	6116401		12-11-2001
				CA	2414290		08-11-2001
				CA	2414296		08-11-2001
				DE	60106423		18-11-2004
				EP	1282643		12-02-2003
				EP	1280820		05-02-2003
				JP	2003531636	Ţ	28-10-2003
				JP	2003531918		28-10-2003
				MO	0183554		08-11-2001
	•			WO US	0183547 2002156000		08-11-2001 24-10-2002
				US	2002156000		20-03-2003
				U.J	E003037333	V.1	20 03-2003